

09/12/017
9/6/00

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Date: 09/06/2000

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Type	L #	Hits	Search Text	DBs	Time Stamp	Comments	Error Definition	Err ors
1	IS&R L1	4106	((("514/8") or ("530/395") or ("530/396") or ("530/399") or ("530/402") or ("530/404") or ("530/411"))).CCLS.	USPAT	2000/09/06 08:29			0
2	BRS L2	6276	lectin or selectin	USPAT	2000/09/06 08:29			0
3	BRS L3	643	l1 and l2	USPAT	2000/09/06 08:34			0
4	BRS L4	616	heparin-binding	USPAT	2000/09/06 08:34			0
5	BRS L5	875	heparin adj binding	USPAT	2000/09/06 08:36			0
6	BRS L6	15508	growth adj factor	USPAT	2000/09/06 08:36			0
7	BRS L7	2698	fibroblast adj l6	USPAT	2000/09/06 08:37			0
8	BRS L8	4245	epidermal adj l6	USPAT	2000/09/06 08:37			0
9	BRS L9	2832	platelet adj derived adj l6	USPAT	2000/09/06 08:38			0
10	BRS L10	6571	l4 or l5 or l7 or l8 or l9	USPAT	2000/09/06 08:38			0
11	BRS L11	824	l1 and l10	USPAT	2000/09/06 08:40			0
12	BRS L12	11427	glycosyls	USPAT	2000/09/06 08:40			0
13	BRS L13	75850	sugar	USPAT	2000/09/06 08:40			0
14	BRS L14	32464	carbohydrate	USPAT	2000/09/06 08:41			0
15	BRS L15	10613	saccharide	USPAT	2000/09/06 08:42			0
16	BRS L16	24699	polysaccharide	USPAT	2000/09/06 08:42			0
17	BRS L17	1847	glycosaminoglycan	USPAT	2000/09/06 08:42			0

	Type	L #	Hits	Search Text	DBs	Time Stamp	Comments	Error Definition	Errors
18	BRS	L18	111803	l12 or l13 or l14 or l15 or l16 or l17	USPAT	2000/09/06 08:45			0
19	BRS	L19	201	l10 with l18	USPAT	2000/09/06 08:45			0
20	BRS	L20	45	l1 and l19	USPAT	2000/09/06 08:46			0

CCXR:
530/395

ORPL:

Settinieri, et al., Characterization of O-Glycosylation Sites in
Recombinant
B-Chain of Platelet-derived Growth Factor Expressed in Yeast
Using Liquid
Secondary Ion Mass Spectrometry, Tandem Mass Spectrometry and
Edman Sequence
Analysis, Biomedican and Environmental Mass Spectrometry, 19:665,
1990.

*proof not
needed*

USPT

US-CL-CURRENT: 435/69.1, 435/69.6, 530/380, 530/402

US-PAT-NO: 6063763

DOCUMENT-IDENTIFIER: US 6063763 A

TITLE: Protease-resistant thrombomodulin analogs

DATE-ISSUED: May 16, 2000

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Light; David Richard	San Mateo	CA	N/A
N/A			
Andrews; William H.	San Mateo	CA	N/A
N/A			
Clarke; Jeffrey Homer	Pacifica	CA	N/A
N/A			
Wydro; Robert Michael	Foster City	CA	N/A
N/A			
Young; Patricia Ann	San Rafael	CA	N/A
N/A			

US-CL-CURRENT: 514/12, 435/69.1, 435/69.6, 530/380, 530/402

ABSTRACT:

The present invention relates to the single-chain thrombomodulin ("TM") and analogs thereof that are not susceptible to cleavage by proteases and retain the biological activity of thrombomodulin, as well as methods of use in, for

example, antithrombotic therapy. Novel proteins, nucleic acid gene sequences, pharmaceuticals and methods of inhibiting thrombotic activity are disclosed.

18 Claims, 5 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 4

DEPL:

259:12246-12251, which is incorporated herein by reference). In Page 1 (DSaunders, 09/06/2000, EAST Version: 1.01.0015)

pel
5,827,824
5,466,668

DEPL: 259:12246-12251, which is incorporated herein by reference). In comparison to native thrombomodulin, preferred TM analogs have been modified to embrace the 6 epidermal growth factor [EGF]-like domains and may also contain the O-linked glycosylation and/or lectin domains.

CCXR:
530/402

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sequence*

USPT

US-CL-CURRENT: 514/2,514/8 , 530/350 , 530/380 , 530/829

US-PAT-NO: 5939390

DOCUMENT-IDENTIFIER: US 5939390 A

TITLE: Pharmaceutical composition

DATE-ISSUED: August 17, 1999

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Flodgaard; Hans	Hellerup	N/A	N/A
DKX			
Rasmussen; Poul Baad	Copenhagen .O	N/A	N/A
DKX			

US-CL-CURRENT: 514/12,514/2 , 514/8 , 530/350 , 530/380 , 530/829

ABSTRACT:

The present invention relates to a pharmaceutical composition for the prevention or treatment of diseases or conditions involving stress injury to cells, the composition comprising

(a) a lipid-containing substance having a lipid portion which is structurally identical with or analogous to a ceramide, conjugated to

(b) a protein capable of binding said lipid-containing substance in such a way that, when the conjugate is contacted with living cells, the lipid-containing substance activates a ceramide-activated protein phosphatase resulting in down-regulation of cellular metabolism, and

(c) a pharmaceutically acceptable diluent or carrier.

31 Claims, 2 Drawing Figures
Exemplary Claim Number: 1 (D. Saunders, 09/06/2000, EAST Version: 1.01.0015)

*probably
not needed*

reducing conditions), the protein being produced in the azurophil granules of polymorphonuclear leukocytes.

CLPR:

2. A composition according to claim 1, wherein the protein to which the lipid-containing substance is conjugated is a heparin-binding protein which, in glycosylated form, has an apparent molecular weight of 28 kD as determined by SDS-PAGE under reducing conditions, the protein being produced in the azurophil granules of polymorphonuclear leukocytes.

CLPV:

(b) a heparin-binding protein which, in glycosylated form, has an apparent molecular weight of 28 kD, the protein being produced in the azurophil granules of polymorphonuclear leukocytes and is capable of binding said lipid-containing substance in such a way that, when the conjugate is contacted with living cells, the lipid-containing substance activates a ceramide-activated protein phosphatase resulting in down-regulation of cellular metabolism.

CLPV:

(b) a heparin-binding protein which, in glycosylated form, has an apparent molecular weight of 28 kD, the protein being produced in the azurophil granules of polymorphonuclear leukocytes and is capable of binding said lipid-containing substance in such a way that, when the conjugate is contacted with living cells, the lipid-containing substance activates a ceramide-activated protein phosphatase resulting in

cells, the lipid-containing substance activates a
ceramide-activated protein
phosphatase resulting in down-regulation of cellular metabolism.

CCXR:
514/8

USPT

US-CL-CURRENT: 514/12,514/21

US-PAT-NO: 5851989

DOCUMENT-IDENTIFIER: US 5851989 A

TITLE: Method of extending the plasma half-life of vascular endothelial cell growth factor

DATE-ISSUED: December 22, 1998

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
Chamow; Steven	San Mateo	CA	N/A
Modi; Nishit	San Bruno	CA	N/A
Schwall; Ralph	Pacifica	CA	N/A
Zioncheck; Thomas	Montara	CA	N/A

US-CL-CURRENT: 514/8,514/12 ,514/21

ABSTRACT:

The invention provides a method for extending the plasma half-life of heparin-binding proteins by coadministering such proteins with a therapeutically acceptable compound capable of inhibiting their binding to a low affinity heparin-like binding site on the surface of cells. In one embodiment of the invention, the heparin-binding protein is a growth factor or selectin. The binding inhibitory compound can, for example, be a purified native heparin preparation, a heparin fragment, or another polyanionic compound, such as dextran sulfate, heparan sulfate, pentosan sulfate, or hyaluronate.

14 Claims, 12 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 7

BSPR:

The present invention is based on the finding that the in vivo half-life of heparin-binding growth factors, such as HGF, in the plasma can be significantly

good

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extended by coadministration with a polyanionic molecule, such as heparin or heparin-derived oligosaccharides. It has further been found that the coadministration of polyanionic molecules increases the amount of heparin-binding proteins entering the plasma after intraperitoneal or subcutaneous administration. Although it is believed that the effect of polyanionic molecules on the bioavailability of heparin-binding proteins is due to their ability to block the binding of heparin-binding proteins to extracellular matrix glycosylaminoglycans, the invention is not limited by this or any other theory in any way. The invention is additionally based on the finding that the presence of polyanionic molecules, and specifically heparin and heparin-like oligosaccharides potentiates the biological activity of heparin-binding proteins (HGF, IL-8) and/or enhances their binding to their respective native receptors (VEGF).

DEPR:

A "functional derivative" of a native heparin-binding protein is a compound that retains at least one qualitative biological activity of the corresponding native protein and has the ability to bind heparin. Functional derivatives include, but are not limited to, fragments of native heparin-binding proteins from any animal species, and derivatives of the native proteins and fragments thereof, wherein the term "derivative" is used to define amino acid sequence and glycosylation variants, and covalent modifications of a native protein, whereas the term "variant" refers to amino acid sequence and glycosylation variants within this definition. An "inhibitor" of a native heparin-binding protein is a compound that inhibits at least one biological activity of the corresponding native protein and has the ability to bind heparin.

DEPR:

The term "glycosylation variant" is used to refer to a heparin-binding protein

molecule having a glycosylation profile different from that of a corresponding native protein. Glycosylation of polypeptides is typically either N-linked or O-linked. N-linked refers to the attachment of the carbohydrate moiety to the side-chain of an asparagine residue. The tripeptide sequences, asparagine-X-serine and asparagine-X-threonine, wherein X is any amino acid except proline, are recognition sequences for enzymatic attachment of the carbohydrate moiety to the asparagine side chain. O-linked glycosylation refers to the attachment of one of the sugars N-acetylgalactosamine, galactose, or xylose to a hydroxyamino acid, most commonly serine or threonine, although 5-hydroxyproline or 5-hydroxylysine may also be involved in O-linked glycosylation. Any difference in the location and/or nature of the carbohydrate moieties present in a glycosylation variant or fragment as compared to its native counterpart is within the scope herein.

DEPR:

The rationale for the use of dextran sulfate is that in a separate series of experiments, we found that premixing of HGF/SF with soluble heparin greatly enhanced its bioavailability from subcutaneous and intraperitoneal sites (unpublished observation). This effect was not specific to heparin and was also observed with other sulfated polysaccharides including pentosan polysulfate, hyaluronate, and dextran sulfate. Dextran sulfate was chosen arbitrarily for the infusion studies. At a 1:10 HGF/SF:dextran sulfate ratio the solution was extremely viscous and probably could not get out of the infusion pump. However, ratios of 1:1 or 1:2 enhanced the effectiveness of HGF/SF infusions. Although the mechanism is not known, we hypothesize that, because HGF/SF binds heparin strongly (Nakamura et al., Proc. Natl. Acad. Sci. USA 83, 6489-6493 [1986]), its absorption from subcutaneous and intraperitoneal sites is low due to trapping by interaction with

heparin
sulfate proteoglycans in the extracellular matrix. We therefore
speculate that
sulfated polysaccharides saturate the heparin binding regions on
HGF/SF,
thereby preventing interaction with matrix components. Heparin
also decreases
the clearance of HGF/SF, and sulfated polysaccharides enhance
hepatocyte
responses to low doses of HGF/SF.

CCOR:
514/8

USPT

US-CL-CURRENT: 435/101,514/12 ,514/21 ,514/54 ,530/399 ,530/412
,530/413
,536/123 ,536/21 ,536/53 ,536/55.3

US-PAT-NO: 5849722

DOCUMENT-IDENTIFIER: US 5849722 A

TITLE: Oligosaccharide having affinity for fibroblast growth factor and

process for producing same

DATE-ISSUED: December 15, 1998

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Habuchi; Hiroko JPX	Aichi	N/A	N/A
Suzuki; Sakaru JPX	Aichi	N/A	N/A
Kimata; Koji JPX	Aichi	N/A	N/A

US-CL-CURRENT: 514/56,435/101 ,514/12 ,514/21 ,514/54 ,530/399
,530/412
,530/413 ,536/123 ,536/21 ,536/53 ,536/55.3

ABSTRACT:

An oligosaccharide having an affinity for fibroblast growth factor, which is composed of 8 to 18 monosaccharide residues, wherein a principal disaccharide unit comprising L-iduronic acid 2-sulfate and N-sulfo-D-glucosamine and a process for producing the oligosaccharide comprising digesting heparan sulfate.

8 Claims, 4 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 4

BSPR:

Page 1 (DSaunders, 09/06/2000, EAST Version: 1.01.0015)
JP-A-2-40399 discloses a complex consisting of α -lvcosaminodlvcan

purified?

application"). This publication describes that the invention was accomplished based on a finding that stability of the FGF mutein increases markedly when glycosaminoglycan such as heparan sulfate and low molecular weight heparan sulfates prepared using hydrogen peroxide is added to an aqueous solution of the FGF mutein.

BSPR:

In addition, JP-A-63-66192 (hereinafter referred to as "Sanofi application") illustrates an invention entitled "Heparin-based Oligosaccharides having Affinity for Cell Growth Factors". The above invention aims at providing heparin type or heparan sulfate type oligosaccharides having markedly high affinities for heparin-binding cell growth factors, which can be obtained, for example, by a process which comprises the steps of: subjecting natural heparin or natural heparan sulfate chain which serves as a starting material to depolymerization (molecular weight reduction) with nitric acid, heparinase, heparitinase or periodic acid; subjecting the resulting product to alcohol precipitation for separating a fraction of saccharides having 10 monosaccharide residues or less and a fraction of saccharides having more than 10 monosaccharide residues; applying the fraction of saccharides having 10 monosaccharide residues or less to an agarose-acrylamide column for separating into a disaccharide fraction, a tetrasaccharide fraction, a hexasaccharide fraction, an octasaccharide fraction and a decasaccharide fraction; and removing chains having no affinity or medium affinity for FGF by

BSPR:

Thus, a complex of the FGF mutein with glycosaminoglycan according to the Sanofi application is composed of the FGF mutein which is not a naturally occurring fibroblast growth factor, because certain amino acids of human basic fibroblast growth factor are replaced with other amino acids. In addition, though it discloses a low molecular weight heparan sulfate as an example of glycosaminoglycan, its illustrative description includes only a complex which consists of the FGF mutein and a relatively long-chained heparin or heparan sulfate. Such a complex possibly might have pharmacologically and physiologically unnecessary or improper structural moieties which react, for example, with antithrombin III, heparin cofactor II, platelet factor 4 and the like.

CCXR:
530/399

USPT

US-CL-CURRENT: 514/2, 514/54, 514/56, 514/57, 514/61, 514/62
514/8, 536/123.1
536/123.13, 536/21, 536/56

US-PAT-NO: 5783568

DOCUMENT-IDENTIFIER: US 5783568 A

TITLE: Methods for treating cancer and other cell proliferative diseases

DATE-ISSUED: July 21, 1998

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
Schlessinger; Joseph	New York	NY	N/A
N/A			
Lax; Irit	Fair Lawn	NJ	N/A
N/A			
Ladbury; John E.	New York	NY	N/A
N/A			
Tang; Peng Cho	Moraga	CA	N/A
N/A			

US-CL-CURRENT: 514/53, 514/2, 514/54, 514/56, 514/57, 514/61
514/62, 514/8
536/123.1, 536/123.13, 536/21, 536/56

ABSTRACT:

The present invention relates to a method of treating in a mammal certain cancers, other cell proliferative diseases, and/or angiogenesis by using a salt or complex of a sulfated saccharide. The invention also relates to the use of mutant heparin binding growth factors that bind to the growth factor receptor, but not to heparin. The invention also provides pharmaceutical compositions for such methods.

9 Claims, 9 Drawing figures
Exemplary Claim Number: (D)saunders, 09/06/2000, EAST Version: 1.01.0015)

Not relevant

dimerization and activation of the receptor involved in the condition treated. Thus, the sulfated compounds inhibit cancer or cell proliferative diseases by inhibiting the activity of heparin-binding growth factors.

DEPR:

The saccharide component of the sulfated saccharide used in accordance with the invention is a monosaccharide, for example, xylose, fructose or glucose, an oligosaccharide, for example, a disaccharide such as sucrose, lactose, maltose or cellobiose, or maltotriose, maltotetraose, maltoheptose, or maltohexose, or fragments of heparin small enough not to bind more than one heparin-binding growth factor at a time or a subunit of any of such saccharides.

CCXR:
514/8

USPT

US-CL-CURRENT: 514/21,530/380 , 530/395

US-PAT-NO: 5814602

DOCUMENT-IDENTIFIER: US 5814602 A

TITLE: Heparin-binding proteins

DATE-ISSUED: September 29, 1998

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Flodgaard; Hans	Hellerup	N/A	N/A
DKX			
Ostergaard; Erik	Vanlose	N/A	N/A
DKX			
Thomsen; Johannes	Kokkedal	N/A	N/A
DKX			
Bayne; Stephen	Roskilde	N/A	N/A
DKX			

US-CL-CURRENT: 514/8,514/21 , 530/380 , 530/395

ABSTRACT:

A heparin-binding protein (HBP) which has, in glycosylated state, an apparent molecular weight of about 28 kDa, determined by SDS-PAGE under reducing conditions, and exhibits angiogenic properties in vivo.
10 Claims, 6 Drawing figures
Exemplary Claim Number: 1
Number of Drawing Sheets: 6

ABPL:

A heparin-binding protein (HBP) which has, in glycosylated state, an apparent molecular weight of about 28 kDa, determined by SDS-PAGE under reducing conditions, and exhibits angiogenic properties in vivo.

CCOR:
514/8

performed

USPT

US-CL-CURRENT: 435/69.1, 530/402

US-PAT-NO: 5686572

DOCUMENT-IDENTIFIER: US 5686572 A

TITLE: Domains of extracellular region of human platelet derived growth factor

receptor polypeptides

DATE-ISSUED: November 11, 1997

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Wolf; David	Palo Alto	CA	N/A
N/A			
Tomlinson; James E.	San Francisco	CA	N/A
N/A			
Fretto; Larry J.	Belmont	CA	N/A
N/A			
Giese; Neill A.	San Francisco	CA	N/A
N/A			
Escobedo; Jaime A.	San Francisco	CA	N/A
N/A			
Williams; Lewis Thomas	Tiburon	CA	N/A
N/A			

US-CL-CURRENT: 530/350, 435/69.1, 530/402

ABSTRACT:

Defined constructs of modified human platelet-derived growth factor receptor

polypeptides are provided. Extracellular region domain structures are

identified and modifications and combinatorial rearrangements of the receptor

segments are provided. Both cell bound and soluble forms of modified segments

are made available, as are methods for assays using them, allowing for

screening of ligand analogues

8 Claims, 4 Drawing figures

rejection PD 6/2
Not relevant

factor receptor polypeptide. Primary structures of two homologous forms of polypeptides have been reported. A type B receptor nucleic acid and its corresponding polypeptide sequence from mouse are reported in Yarden et al. (1986) Nature 323: 226-232; and a homologous genetic sequence has been isolated from humans. See application Ser. No. 07/771,829 which is a continuation of Ser. No. 07/309,332, now abandoned. A human type A receptor sequence is reported in Matsui et al. (1989) Science 243: 800-803. Although the two different forms of the receptor polypeptides are homologous, they are encoded by two separate genes.

CCXR:
530/402

ORPL:
Daniel et al. (1987) "Biosynthetic and Glycosylation Studies of Cell Surface Platelet-Derived Growth Factor Receptors" J. Biol. Chem. 262:9778-9784.

ORPL:
Keating et al. (1989) "Platelet-Derived Growth Factor Receptor Inducibility Is Acquired Immediately after Translation and Does Not Require Glycosylation" J. Biol. Chem. 264:9129-9132.

USPT

US-CL-CURRENT: 435/69.1, 514/21, 530/350, 530/389.2, 530/399

US-PAT-NO: 5686415

DOCUMENT-IDENTIFIER: US 5686415 A

TITLE: Method for the treatment of colon epithelial cells in vivo

DATE-ISSUED: November 11, 1997

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Carnahan; Josette	Newbury Park	CA	N/A
N/A			
Fran.cedilla.oise	Thousand Oaks	CA	N/A
N/A			
Hara; Shinichi	Thousand Oaks	CA	N/A
N/A			
Lu; Hsieng Sen	Boulder	CO	N/A
N/A			
Mayer; John Philip	Thousand Oaks	CA	N/A
N/A			

Yoshinaga; Steven
Kiyoshi

US-CL-CURRENT: 514/12, 435/69.1, 514/21, 530/350, 530/389.2, 530/399

ABSTRACT:

Colon epithelial cells are stimulated to multiply, grow and mature by contacting them in vivo with peptides derived from the EGF-like domain of

proteins of the NDF/herregulin family.

5 Claims, 8 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 5

BSPR:

The cell membrane-bound precursor for these proteins (referred to in this document) is disclosed in US 5,000,000, 5,000,001, 5,000,002, 5,000,003, 5,000,004, 5,000,005, 5,000,006, 5,000,007, 5,000,008, 5,000,009, 5,000,010, 5,000,011, 5,000,012, 5,000,013, 5,000,014, 5,000,015, 5,000,016, 5,000,017, 5,000,018, 5,000,019, 5,000,020, 5,000,021, 5,000,022, 5,000,023, 5,000,024, 5,000,025, 5,000,026, 5,000,027, 5,000,028, 5,000,029, 5,000,030, 5,000,031, 5,000,032, 5,000,033, 5,000,034, 5,000,035, 5,000,036, 5,000,037, 5,000,038, 5,000,039, 5,000,040, 5,000,041, 5,000,042, 5,000,043, 5,000,044, 5,000,045, 5,000,046, 5,000,047, 5,000,048, 5,000,049, 5,000,050, 5,000,051, 5,000,052, 5,000,053, 5,000,054, 5,000,055, 5,000,056, 5,000,057, 5,000,058, 5,000,059, 5,000,060, 5,000,061, 5,000,062, 5,000,063, 5,000,064, 5,000,065, 5,000,066, 5,000,067, 5,000,068, 5,000,069, 5,000,070, 5,000,071, 5,000,072, 5,000,073, 5,000,074, 5,000,075, 5,000,076, 5,000,077, 5,000,078, 5,000,079, 5,000,080, 5,000,081, 5,000,082, 5,000,083, 5,000,084, 5,000,085, 5,000,086, 5,000,087, 5,000,088, 5,000,089, 5,000,090, 5,000,091, 5,000,092, 5,000,093, 5,000,094, 5,000,095, 5,000,096, 5,000,097, 5,000,098, 5,000,099, 5,000,100, 5,000,101, 5,000,102, 5,000,103, 5,000,104, 5,000,105, 5,000,106, 5,000,107, 5,000,108, 5,000,109, 5,000,110, 5,000,111, 5,000,112, 5,000,113, 5,000,114, 5,000,115, 5,000,116, 5,000,117, 5,000,118, 5,000,119, 5,000,120, 5,000,121, 5,000,122, 5,000,123, 5,000,124, 5,000,125, 5,000,126, 5,000,127, 5,000,128, 5,000,129, 5,000,130, 5,000,131, 5,000,132, 5,000,133, 5,000,134, 5,000,135, 5,000,136, 5,000,137, 5,000,138, 5,000,139, 5,000,140, 5,000,141, 5,000,142, 5,000,143, 5,000,144, 5,000,145, 5,000,146, 5,000,147, 5,000,148, 5,000,149, 5,000,150, 5,000,151, 5,000,152, 5,000,153, 5,000,154, 5,000,155, 5,000,156, 5,000,157, 5,000,158, 5,000,159, 5,000,160, 5,000,161, 5,000,162, 5,000,163, 5,000,164, 5,000,165, 5,000,166, 5,000,167, 5,000,168, 5,000,169, 5,000,170, 5,000,171, 5,000,172, 5,000,173, 5,000,174, 5,000,175, 5,000,176, 5,000,177, 5,000,178, 5,000,179, 5,000,180, 5,000,181, 5,000,182, 5,000,183, 5,000,184, 5,000,185, 5,000,186, 5,000,187, 5,000,188, 5,000,189, 5,000,190, 5,000,191, 5,000,192, 5,000,193, 5,000,194, 5,000,195, 5,000,196, 5,000,197, 5,000,198, 5,000,199, 5,000,200, 5,000,201, 5,000,202, 5,000,203, 5,000,204, 5,000,205, 5,000,206, 5,000,207, 5,000,208, 5,000,209, 5,000,210, 5,000,211, 5,000,212, 5,000,213, 5,000,214, 5,000,215, 5,000,216, 5,000,217, 5,000,218, 5,000,219, 5,000,220, 5,000,221, 5,000,222, 5,000,223, 5,000,224, 5,000,225, 5,000,226, 5,000,227, 5,000,228, 5,000,229, 5,000,230, 5,000,231, 5,000,232, 5,000,233, 5,000,234, 5,000,235, 5,000,236, 5,000,237, 5,000,238, 5,000,239, 5,000,240, 5,000,241, 5,000,242, 5,000,243, 5,000,244, 5,000,245, 5,000,246, 5,000,247, 5,000,248, 5,000,249, 5,000,250, 5,000,251, 5,000,252, 5,000,253, 5,000,254, 5,000,255, 5,000,256, 5,000,257, 5,000,258, 5,000,259, 5,000,260, 5,000,261, 5,000,262, 5,000,263, 5,000,264, 5,000,265, 5,000,266, 5,000,267, 5,000,268, 5,000,269, 5,000,270, 5,000,271, 5,000,272, 5,000,273, 5,000,274, 5,000,275, 5,000,276, 5,000,277, 5,000,278, 5,000,279, 5,000,280, 5,000,281, 5,000,282, 5,000,283, 5,000,284, 5,000,285, 5,000,286, 5,000,287, 5,000,288, 5,000,289, 5,000,290, 5,000,291, 5,000,292, 5,000,293, 5,000,294, 5,000,295, 5,000,296, 5,000,297, 5,000,298, 5,000,299, 5,000,300, 5,000,301, 5,000,302, 5,000,303, 5,000,304, 5,000,305, 5,000,306, 5,000,307, 5,000,308, 5,000,309, 5,000,310, 5,000,311, 5,000,312, 5,000,313, 5,000,314, 5,000,315, 5,000,316, 5,000,317, 5,000,318, 5,000,319, 5,000,320, 5,000,321, 5,000,322, 5,000,323, 5,000,324, 5,000,325, 5,000,326, 5,000,327, 5,000,328, 5,000,329, 5,000,330, 5,000,331, 5,000,332, 5,000,333, 5,000,334, 5,000,335, 5,000,336, 5,000,337, 5,000,338, 5,000,339, 5,000,340, 5,000,341, 5,000,342, 5,000,343, 5,000,344, 5,000,345, 5,000,346, 5,000,347, 5,000,348, 5,000,349, 5,000,350, 5,000,351, 5,000,352, 5,000,353, 5,000,354, 5,000,355, 5,000,356, 5,000,357, 5,000,358, 5,000,359, 5,000,360, 5,000,361, 5,000,362, 5,000,363, 5,000,364, 5,000,365, 5,000,366, 5,000,367, 5,000,368, 5,000,369, 5,000,370, 5,000,371, 5,000,372, 5,000,373, 5,000,374, 5,000,375, 5,000,376, 5,000,377, 5,000,378, 5,000,379, 5,000,380, 5,000,381, 5,000,382, 5,000,383, 5,000,384, 5,000,385, 5,000,386, 5,000,387, 5,000,388, 5,000,389, 5,000,390, 5,000,391, 5,000,392, 5,000,393, 5,000,394, 5,000,395, 5,000,396, 5,000,397, 5,000,398, 5,000,399, 5,000,400, 5,000,401, 5,000,402, 5,000,403, 5,000,404, 5,000,405, 5,000,406, 5,000,407, 5,000,408, 5,000,409, 5,000,410, 5,000,411, 5,000,412, 5,000,413, 5,000,414, 5,000,415, 5,000,416, 5,000,417, 5,000,418, 5,000,419, 5,000,420, 5,000,421, 5,000,422, 5,000,423, 5,000,424, 5,000,425, 5,000,426, 5,000,427, 5,000,428, 5,000,429, 5,000,430, 5,000,431, 5,000,432, 5,000,433, 5,000,434, 5,000,435, 5,000,436, 5,000,437, 5,000,438, 5,000,439, 5,000,440, 5,000,441, 5,000,442, 5,000,443, 5,000,444, 5,000,445, 5,000,446, 5,000,447, 5,000,448, 5,000,449, 5,000,450, 5,000,451, 5,000,452, 5,000,453, 5,000,454, 5,000,455, 5,000,456, 5,000,457, 5,000,458, 5,000,459, 5,000,460, 5,000,461, 5,000,462, 5,000,463, 5,000,464, 5,000,465, 5,000,466, 5,000,467, 5,000,468, 5,000,469, 5,000,470, 5,000,471, 5,000,472, 5,000,473, 5,000,474, 5,000,475, 5,000,476, 5,000,477, 5,000,478, 5,000,479, 5,000,480, 5,000,481, 5,000,482, 5,000,483, 5,000,484, 5,000,485, 5,000,486, 5,000,487, 5,000,488, 5,000,489, 5,000,490, 5,000,491, 5,000,492, 5,000,493, 5,000,494, 5,000,495, 5,000,496, 5,000,497, 5,000,498, 5,000,499, 5,000,500, 5,000,501, 5,000,502, 5,000,503, 5,000,504, 5,000,505, 5,000,506, 5,000,507, 5,000,508, 5,000,509, 5,000,510, 5,000,511, 5,000,512, 5,000,513, 5,000,514, 5,000,515, 5,000,516, 5,000,517, 5,000,518, 5,000,519, 5,000,520, 5,000,521, 5,000,522, 5,000,523, 5,000,524, 5,000,525, 5,000,526, 5,000,527, 5,000,528, 5,000,529, 5,000,530, 5,000,531, 5,000,532, 5,000,533, 5,000,534, 5,000,535, 5,000,536, 5,000,537, 5,000,538, 5,000,539, 5,000,540, 5,000,541, 5,000,542, 5,000,543, 5,000,544, 5,000,545, 5,000,546, 5,000,547, 5,000,548, 5,000,549, 5,000,550, 5,000,551, 5,000,552, 5,000,553, 5,000,554, 5,000,555, 5,000,556, 5,000,557, 5,000,558, 5,000,559, 5,000,560, 5,000,561, 5,000,562, 5,000,563, 5,000,564, 5,000,565, 5,000,566, 5,000,567, 5,000,568, 5,000,569, 5,000,570, 5,000,571, 5,000,572, 5,000,573, 5,000,574, 5,000,575, 5,000,576, 5,000,577, 5,000,578, 5,000,579, 5,000,580, 5,000,581, 5,000,582, 5,000,583, 5,000,584, 5,000,585, 5,000,586, 5,000,587, 5,000,588, 5,000,589, 5,000,590, 5,000,591, 5,000,592, 5,000,593, 5,000,594, 5,000,595, 5,000,596, 5,000,597, 5,000,598, 5,000,599, 5,000,600, 5,000,601, 5,000,602, 5,000,603, 5,000,604, 5,000,605, 5,000,606, 5,000,607, 5,000,608, 5,000,609, 5,000,610, 5,000,611, 5,000,612, 5,000,613, 5,000,614, 5,000,615, 5,000,616, 5,000,617, 5,000,618, 5,000,619, 5,000,620, 5,000,621, 5,000,622, 5,000,623, 5,000,624, 5,000,625, 5,000,626, 5,000,627, 5,000,628, 5,000,629, 5,000,630, 5,000,631, 5,000,632, 5,000,633, 5,000,634, 5,000,635, 5,000,636, 5,000,637, 5,000,638, 5,000,639, 5,000,640, 5,000,641, 5,000,642, 5,000,643, 5,000,644, 5,000,645, 5,000,646, 5,000,647, 5,000,648, 5,000,649, 5,000,650, 5,000,651, 5,000,652, 5,000,653, 5,000,654, 5,000,655, 5,000,656, 5,000,657, 5,000,658, 5,000,659, 5,000,660, 5,000,661, 5,000,662, 5,000,663, 5,000,664, 5,000,665, 5,000,666, 5,000,667, 5,000,668, 5,000,669, 5,000,670, 5,000,671, 5,000,672, 5,000,673, 5,000,674, 5,000,675, 5,000,676, 5,000,677, 5,000,678, 5,000,679, 5,000,680, 5,000,681, 5,000,682, 5,000,683, 5,000,684, 5,000,685, 5,000,686, 5,000,687, 5,000,688, 5,000,689, 5,000,690, 5,000,691, 5,000,692, 5,000,693, 5,000,694, 5,000,695, 5,000,696, 5,000,697, 5,000,698, 5,000,699, 5,000,700, 5,000,701, 5,000,702, 5,000,703, 5,000,704, 5,000,705, 5,000,706, 5,000,707, 5,000,708, 5,000,709, 5,000,710, 5,000,711, 5,000,712, 5,000,713, 5,000,714, 5,000,715, 5,000,716, 5,000,717, 5,000,718, 5,000,719, 5,000,720, 5,000,721, 5,000,722, 5,000,723, 5,000,724, 5,000,725, 5,000,726, 5,000,727, 5,000,728, 5,000,729, 5,000,730, 5,000,731, 5,000,732, 5,000,733, 5,000,734, 5,000,735, 5,000,736, 5,000,737, 5,000,738, 5,000,739, 5,000,740, 5,000,741, 5,000,742, 5,000,743, 5,000,744, 5,000,745, 5,000,746, 5,000,747, 5,000,748, 5,000,749, 5,000,750, 5,000,751, 5,000,752, 5,000,753, 5,000,754, 5,000,755, 5,000,756, 5,000,757, 5,000,758, 5,000,759, 5,000,760, 5,000,761, 5,000,762, 5,000,763, 5,000,764, 5,000,765, 5,000,766, 5,000,767, 5,000,768, 5,000,769, 5,000,770, 5,000,771, 5,000,772, 5,000,773, 5,000,774, 5,000,775, 5,000,776, 5,000,777, 5,000,778, 5,000,779, 5,000,780, 5,000,781, 5,000,782, 5,000,783, 5,000,784, 5,000,785, 5,000,786, 5,000,787, 5,000,788, 5,000,789, 5,000,790, 5,000,791, 5,000,792, 5,000,793, 5,000,794, 5,000,795, 5,000,796, 5,000,797, 5,000,798, 5,000,799, 5,000,800, 5,000,801, 5,000,802, 5,000,803, 5,000,804, 5,000,805, 5,000,806, 5,000,807, 5,000,808, 5,000,809, 5,000,810, 5,000,811, 5,000,812, 5,000,813, 5,000,814, 5,000,815, 5,000,816, 5,000,817, 5,000,818, 5,000,819, 5,000,820, 5,000,821, 5,000,822, 5,000,823, 5,000,824, 5,000,825, 5,000,826, 5,000,827, 5,000,828, 5,000,829, 5,000,830, 5,000,831, 5,000,832, 5,000,833, 5,000,834, 5,000,835, 5,000,836, 5,000,837, 5,000,838, 5,000,839, 5,000,840, 5,000,841, 5,000,842, 5,000,843, 5,000,844, 5,000,845, 5,000,846, 5,000,847, 5,000,848, 5,000,849, 5,000,850, 5,000,851, 5,000,852, 5,000,853, 5,000,854, 5,000,855, 5,000,856, 5,000,857, 5,000,858, 5,000,859, 5,000,860, 5,000,861, 5,000,862, 5,000,863, 5,000,864, 5,000,865, 5,000,866, 5,000,867, 5,000,868, 5,000,869, 5,000,870, 5,000,871, 5,000,872, 5,000,873, 5,000,874, 5,000,875, 5,000,876, 5,000,877, 5,000,878, 5,000,879, 5,000,880, 5,000,881, 5,000,882, 5,000,883, 5,000,884, 5,000,885, 5,000,886, 5,000,887, 5,000,888, 5,000,889, 5,000,890, 5,000,891, 5,000,892, 5,000,893, 5,000,894, 5,000,895, 5,000,896, 5,000,897, 5,000,898, 5,000,899, 5,000,900, 5,000,901, 5,000,902, 5,000,903, 5,000,904, 5,000,905, 5,000,906, 5,000,907, 5,000,908, 5,000,909, 5,000,910, 5,000,911, 5,000,912, 5,000,913, 5,000,914, 5,000,915, 5,000,916, 5,000,917, 5,000,918, 5,000,919, 5,000,920, 5,000,921, 5,000,922, 5,000,923, 5,000,924, 5,000,925, 5,000,926, 5,000,927, 5,000,928, 5,000,929, 5,000,930, 5,000,931, 5,000,932, 5,000,933, 5,000,934, 5,000,935, 5,000,936, 5,000,937, 5,000,938, 5,000,939, 5,000,940, 5,000,941, 5,000,942, 5,000,943, 5,000,944, 5,000,945, 5,000,946, 5,000,947, 5,000,948, 5,000,949, 5,000,950, 5,000,951, 5,000,952, 5,000,953, 5,000,954, 5,000,955, 5,000,956, 5,000,957, 5,000,958, 5,000,959, 5,000,960, 5,000,961, 5,000,962, 5,000,963, 5,000,964, 5,000,965, 5,000,966, 5,000,967, 5,000,968, 5,000,969, 5,000,970, 5,000,971, 5,000,972, 5,000,973, 5,000,974, 5,000,975, 5,000,976, 5,000,977, 5,000,978, 5,000,979, 5,000,980, 5,000,981, 5,000,982, 5,000,983, 5,000,984, 5,000,985, 5,000,986, 5,000,987, 5,000,988, 5,000,989, 5,000,990, 5,000,991, 5,000,992, 5,000,993, 5,000,994, 5,000,995, 5,000,996, 5,000,997, 5,000,998, 5,000,999, 5,000,1000, 5,000,1001, 5,000,1002, 5,000,1003, 5,000,1004, 5,000,1005, 5,000,1006, 5,000,1007, 5,000,1008, 5,000,1009, 5,000,1010, 5,000,1011, 5,000,1012, 5,000,1013, 5,000,1014, 5,000,101

immunoglobulin(Ig)-like domain (approximately 70 amino acid residues), a so-called "spacer" domain that contains multiple binding sites for N- and O-linked glycosylation, an epidermal growth factor (EGF)-like domain of about 60-75 amino acid residues that includes 6 cysteine residues, a hydrophobic region of about 25 amino acid residues that functions as a transmembrane domain, and a "cytoplasmic tail" which can vary in length. Some of these transmembranous precursor forms undergo proteolytic cleavage in the cell at both the N-terminus and at the short stretch of sequence (juxtamembrane) that connects the EGF-like domain with the transmembrane domain. Depending on the amino acid sequence in this juxtamembrane region, the NDF/hereginins have been designated subtype 1, subtype 2, subtype 3, etc. Additional variations comprise two forms of the C-terminal loop of the EGF-like domain, which are termed alpha(.alpha.) and beta(.beta.), depending on the amino acid sequence in this region; Wen et al., Molecular and Cellular Biology, Volume 14, Number 3, pages 1909-1919 (1994).

DRPR:

FIG. 1. This figure is a schematic drawing (not in proportional scale) of the extracellular structure of human NDF/hereginin, comprising (distal from the end that binds to the cell membrane): the putative N-terminal "heparin-binding" region, an immunoglobulin (Ig)-like domain, a carbohydrate (or "spacer") domain, and an EGF-like domain proximal to the C-terminal end. The open circles in the spacer domain represent O-linked sugars and the

towards the C-terminal end. The EGF-like domain begins approximately at amino acid residue position 177 (marked) and ends approximately at amino acid residue position 228 (not marked).

CCXR:
530/399

USPT

US-CL-CURRENT: 435/252.3, 435/252.33 , 435/255.1 , 435/320.1 , 435/69.1 , 435/69.4 , 530/399 , 536/23.51

US-PAT-NO: 5464943

DOCUMENT-IDENTIFIER: US 5464943 A

TITLE: DNA encoding glycosylated FGF and production thereof

DATE-ISSUED: November 7, 1995

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Senoo; Masaharu JPX	Toyonaka	N/A	N/A
Sasada; Reiko JPX	Kyoto	N/A	N/A
Igarashi; Koichi JPX	Kyoto	N/A	N/A

US-CL-CURRENT: 536/23.5, 435/252.3 , 435/252.33 , 435/255.1 , 435/320.1 , 435/69.1 , 435/69.4 , 530/399 , 536/23.51

ABSTRACT:

Disclosed are (1) a mutein of a fibroblast growth factor (FGF), the DNA having introduced therein at least one nucleotide sequence coding for a glycosylation site, (2) a DNA coding for the mutein of (1), (3) a vector containing the DNA of (2), (4) a transformant transformed with the vector of (3), and (5) a process for producing the mutein which comprises cultivating in a culture medium the transformant of a yeast or animal cell transformed with a vector of (3), and producing and accumulating the mutein of (1) in the culture medium, whereby the FGF mutein is produced.
Page 1 of 1
USPT, 09/06/2000, EAST Version: 1.01.0015)

the DNA having introduced therein at least one nucleotide sequence coding for a glycosylation site, (2) a DNA coding for the mutein of (1), (3) a vector containing the DNA of (2), (4) a transformant transformed with the vector of (3), and (5) a process for producing the mutein which comprises cultivating in a culture medium the transformant of a yeast or animal cell transformed with a vector of (3), and producing and accumulating the mutein of (1) in the culture medium, whereby the FGF mutein into which at least one glycosylation site has been introduced is improved in productivity, stability and activities, and advantageously used as medicine.

BSPV:

(1) a mutein of a fibroblast growth factor (FGF) into which at least one glycosylation site has been introduced,

CLPR:

1. A DNA coding for a mutein of a naturally occurring fibroblast growth factor (FGF), the DNA having artificially introduced therein at least one nucleotide sequence coding for a glycosylation site which is represented by -Asn-X-Y, wherein X is Gly, Lys, Val or Ala; Y is Thr, Ser or Cys; subject to the limitation that -X-Y- is not -Gly-Ser-.

CLPR:

3. A plasmid containing a DNA coding for a mutein of a naturally occurring fibroblast growth factor (FGF), the DNA having artificially introduced therein at least one nucleotide sequence coding for a glycosylation site which is

artificially introduced at least one nucleotide sequence coding for a glycosylation site which is represented by -Asn-X-Y-, wherein X is Gly, Lys, Val or Ala; Y is Thr, Ser or Cys; subject to the limitation that -X-Y- is not -Gly-Ser-.

CLPR:

9. A process for producing a mutein of a naturally occurring fibroblast growth factor (FGF), into which has been artificially introduced at least one glycosylation site which is represented by -Asn-X-Y-, wherein X is Gly, Lys, Val or Ala; Y is Thr, Ser or Cys; subject to the limitation that -X-Y- is not -Gly-Ser- which comprises;

CLPV:

cultivating in a culture medium a yeast or animal cell transformant transformed with a vector containing a DNA coding for a mutein of a naturally occurring fibroblast growth factor (FGF), the DNA having introduced therein at least one nucleotide sequence coding for a glycosylation site which is represented by -Asn-X-Y-, wherein X is Gly, Lys, Val or Ala; Y is Thr, Ser or Cys; subject to the limitation that -X-Y- is not -Gly-Ser-, and recovering said mutein from the culture medium.

CCXR:

530/399

USPT

US-CL-CURRENT: 435/69.1, 435/69.4 , 435/69.5 , 530/397

US-PAT-NO: 5360896

DOCUMENT-IDENTIFIER: US 5360896 A

TITLE: Glycosylated FGF

DATE-ISSUED: November 1, 1994

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Senoo; Masaharu JPX	Osaka	N/A	N/A
Sasada; Reiko JPX	Kyoto	N/A	N/A
Igarashi; Koichi JPX	Kyoto	N/A	N/A

US-CL-CURRENT: 530/399, 435/69.1 , 435/69.4 , 435/69.5 , 530/397

ABSTRACT:

Disclosed are (1) a mutein of a fibroblast growth factor (FGF), the DNA having introduced therein at least one nucleotide sequence coding for a glycosylation site, (2) a DNA coding for the mutein of (1), (3) a vector containing the DNA of (2), (4) a transformant transformed with the vector of (3), and (5) a process for producing the mutein which comprises cultivating in a culture medium the transformant of a yeast or animal cell transformed with a vector of (3), and producing and accumulating the mutein of (1) in the culture medium, whereby the FGF mutein into which at least one glycosylation site has been introduced is improved in productivity, stability and activities, and.

advantageously used as a medicine.
16 Claims, 19 Drawing Figures
D. Saunders, 09/06/2000, EAST Version: 1.01.0015)

containing the DNA of (2), (4) a transformant transformed with the vector of (3), and (5) a process for producing the mutein which comprises cultivating in a culture medium the transformant of a yeast or animal cell transformed with a vector of (3), and producing and accumulating the mutein of (1) in the culture medium, whereby the FGF mutein into which at least one glycosylation site has been introduced is improved in productivity, stability and activities, and advantageously used as medicine.

BSPV:

(1) a mutein of a fibroblast growth factor (FGF) into which at least one glycosylation site has been introduced,

CLPR:

1. A mutein of a naturally occurring fibroblast growth factor (FGF) into which has been artificially introduced at least one glycosylation site which is represented by -Asn-X-Y-, wherein X is Gly, Lys, Val or Ala; Y is Thr, Ser or Cys; subject to the limitation that -X-Y- is not -Gly-Ser-.

CLPR:

12. The mutein according to claim 1, wherein the mutein is produced from a DNA coding for a mutein of a fibroblast growth factor (FGF), the DNA having artificially introduced therein at least one nucleotide sequence coding for the glycosylation site.

CCOR:

530/399

USPT

US-CL-CURRENT: 514/54,514/56 ,514/59 ,514/885 ,514/886 ,514/889 ,530/399

US-PAT-NO: 5288704

DOCUMENT-IDENTIFIER: US 5288704 A

TITLE: Synergistic composition comprising a fibroblast growth factor and a

sulfated polysaccharide, for use as antiviral agent

DATE-ISSUED: February 22, 1994

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE
COUNTRY			
Ungheri; Domenico ITX	Parabiago	N/A	N/A
Garofano; Luisa ITX	Monza	N/A	N/A
Battistini; Carlo ITX	Novate Milanese	N/A	N/A
Carminati; Paolo ITX	Milan	N/A	N/A
Mazue; Guy ITX	Milan	N/A	N/A

US-CL-CURRENT: 514/12,514/54 ,514/56 ,514/59 ,514/885 ,514/886 ,514/889 ,530/399

ABSTRACT:

A pharmaceutical composition is provided for use in the prevention or treatment of viral infections caused by enveloped viruses. The composition comprises a fibroblast growth factor, a sulfated polysaccharide with antiviral activity, and one or more pharmaceutically acceptable carriers. The fibroblast growth factor may be a basic fibroblast growth factor or an analogue thereof, page 1 (Dsaunders, 09/06/2000, EAST Version: 1.01.0015) and the polysaccharide may be a carrageenan, heparin, dextran

Synergistic composition comprising a fibroblast growth factor and a sulfated polysaccharide, for use as antiviral agent

ABPL:

A pharmaceutical composition is provided for use in the prevention or treatment of viral infections caused by enveloped viruses. The composition comprises a fibroblast growth factor, a sulfated polysaccharide with antiviral activity, and one or more pharmaceutically acceptable carriers. The fibroblast growth factor may be a basic fibroblast growth factor or an analogue thereof, and the polysaccharide may be a carrageenan, heparin, dextran sulfate, pentosan polysulfate or a sulfated polysaccharides produced by marine algae belonging to the class of Rhodophyceae.

BSPR:

The present invention relates to a synergistic pharmaceutical composition having antiviral activity which comprises a fibroblast growth factor, a sulfated polysaccharide with antiviral activity and any acceptable pharmaceutical excipient or excipients, for the use in the prevention or treatment of viral infections caused by enveloped viruses.

BSPR:

We have surprisingly found that combining a fibroblast growth factor and a sulfated polysaccharide, the obtained antiviral activity is superior to that expected from the sum of the antiviral activities of the individual constituents of the combination, thus indicating the presence of a synergistic effect. The combination of the two components is therefore more

acceptable pharmaceutical excipient or excipients, for use in the prevention or treatment of viral infections caused by enveloped viruses.

DRPR:

The invention further concerns a process for preparing the above mentioned composition, comprising the step of combining a fibroblast growth factor, and a sulfated polysaccharide in a pharmaceutically acceptable excipient or excipients.

CCXR:

530/399